

The Listing of Claims

1. (Currently Amended) A nucleic acid expression cassette capable of expressing human Factor IX, wherein the cassette is predominantly expressed in the mammalian liver of a post-natal subject, said cassette comprising:

- (a) [[an]] a hepatic locus control element consisting of SEQ ID NO:4 or SEQ ID NO:9;
- (b) [[an]] a heterologous hepatic promoter located 3' to the hepatic locus control element, said promoter consisting of a human α -1 antitrypsin promoter (SEQ ID NO:5);
- (c) a Factor IX coding sequence located 3' to the hepatic promoter, said coding sequence comprising SEQ ID NO:2;
- (d) a polyadenylation signal located 3' to the intron sequence, said polyadenylation signal consisting of SEQ ID NO:6; and
- (e) an intron located 3' to the hepatic promoter and 5' to the polyadenylation signal, wherein said intron consists of SEQ ID NO:1,

wherein elements (a), (b), (c), (d) and (e) are operably linked to express the polypeptide encoded by the coding sequence.

2. (Currently Amended) The expression cassette of claim 1, wherein said cassette directs expression of a therapeutic amount of Factor IX in liver cells for a period of at least 100 days, and further wherein said cassette is expressed in a transgenic animal or as a component of a recombinant adeno-associated virus (AAV) vector.

3. (Currently Amended) The expression cassette of claim 1, wherein said cassette directs expression of a therapeutic amount of the polypeptide in liver cells for a period of at least 300 days, and further wherein said cassette is expressed in a transgenic animal or as a component of a recombinant adeno-associated virus (AAV) vector.

4. (Currently Amended) The expression cassette of claim 1, wherein said cassette directs expression of a therapeutic amount of the polypeptide in liver cells for a period of at least 500 days, and further wherein said cassette is expressed in a transgenic animal or as a component of a recombinant adeno-

associated virus (AAV) vector.

5-14 (Canceled).

15. (Currently Amended) ~~The expression cassette of Claim 1, wherein the cassette further comprises~~
A nucleic acid expression cassette capable of expressing human Factor IX, wherein the cassette is
predominantly expressed in the mammalian liver, said cassette comprising:

_____ (a) a hepatic locus control element consisting of SEQ ID NO:4 or SEQ ID NO:9;

_____ (b) a hepatic promoter located 3' to the hepatic locus control element, said promoter
consisting of SEQ ID NO:5;

_____ (c) a Factor IX coding sequence located 3' to the hepatic promoter, said coding sequence
comprising SEQ ID NO:2;

_____ (d) a polyadenylation signal located 3' to the intron sequence, said polyadenylation signal
consisting of SEQ ID NO:6; and

_____ (e) an intron located 3' to the hepatic promoter and 5' to the polyadenylation signal, wherein
said intron consists of SEQ ID NO:1, and

_____ (f) an untranslated region located 3' to the coding region and the intron, said untranslated
region consisting of SEQ ID NO:7;

_____ wherein elements (a), (b), (c), (d) and (e) are operably linked to express the polypeptide
encoded by the coding sequence.

16. (Previously Presented) The expression cassette of claim 1, wherein the coding sequence encodes
a Factor IX polypeptide consisting of the amino acid sequence set forth in SEQ ID NO:3.

17-23. (Canceled).

24. (Currently Amended) ~~The expression cassette of claim 1, further comprising~~ A nucleic acid
expression cassette capable of expressing human Factor IX, wherein the cassette is predominantly
expressed in the mammalian liver, said cassette comprising:

_____ (a) a hepatic locus control element comprising an enhancer sequence consisting of SEQ ID

NO:8;

(b) a heterologous hepatic promoter located 3' to the hepatic locus control element, said promoter consisting of a human α -1 antitrypsin promoter (SEQ ID NO:5);

(c) a Factor IX coding sequence located 3' to the hepatic promoter, said coding sequence comprising SEQ ID NO:2;

(d) a polyadenylation signal located 3' to the intron sequence, said polyadenylation signal consisting of SEQ ID NO:6; and

(e) an intron located 3' to the hepatic promoter and 5' to the polyadenylation signal, wherein said intron consists of SEQ ID NO:1;

wherein elements (a), (b), (c), (d) and (e) are operably linked to express the polypeptide encoded by the coding sequence.

25-35. (Canceled).

36. (New) The expression cassette of claim 15, wherein said cassette directs expression of a therapeutic amount of Factor IX in liver cells for a period of at least 100 days, and further wherein said cassette is expressed in a transgenic animal or as a component of a recombinant adeno-associated virus (AAV) vector.

37. (New) The expression cassette of claim 15, wherein said cassette directs expression of a therapeutic amount of the polypeptide in liver cells for a period of at least 300 days, and further wherein said cassette is expressed in a transgenic animal or as a component of a recombinant adeno-associated virus (AAV) vector.

38. (New) The expression cassette of claim 15, wherein said cassette directs expression of a therapeutic amount of the polypeptide in liver cells for a period of at least 500 days, and further wherein said cassette is expressed in a transgenic animal or as a component of a combinant adeno-associated virus (AAV) vector.

39. (New) The expression cassette of claim 15, wherein the coding sequence encodes a Factor IX polypeptide consisting of the amino acid sequence set forth in SEQ ID NO:3.

40. (New) The expression cassette of claim 24, wherein said cassette directs expression of a therapeutic amount of Factor IX in liver cells for a period of at least 100 days, and further wherein said cassette is expressed in a transgenic animal or as a component of a recombinant adeno-associated virus (AAV) vector.

41. (New) The expression cassette of claim 24, wherein said cassette directs expression of a therapeutic amount of the polypeptide in liver cells for a period of at least 300 days, and further wherein said cassette is expressed in a transgenic animal or as a component of a recombinant adeno-associated virus (AAV) vector.

42. (New) The expression cassette of claim 24, wherein said cassette directs expression of a therapeutic amount of the polypeptide in liver cells for a period of at least 500 days, and further wherein said cassette is expressed in a transgenic animal or as a component of a recombinant adeno-associated virus (AAV) vector.

43. (New) The expression cassette of claim 24, wherein the coding sequence encodes a Factor IX polypeptide consisting of the amino acid sequence set forth in SEQ ID NO:3.